

FIGURE 1
Induction of bone marrow stem cell proliferation

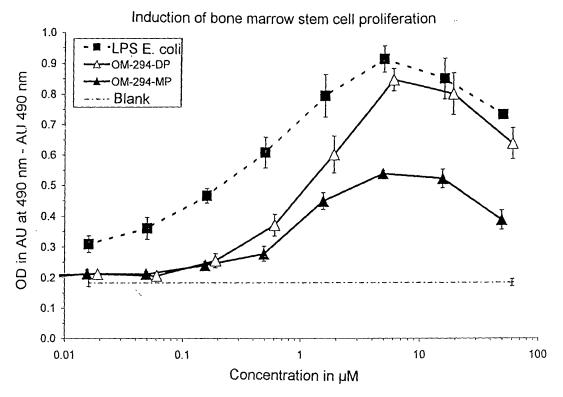


FIGURE 2

Induction of NO production in murine macrophage cells

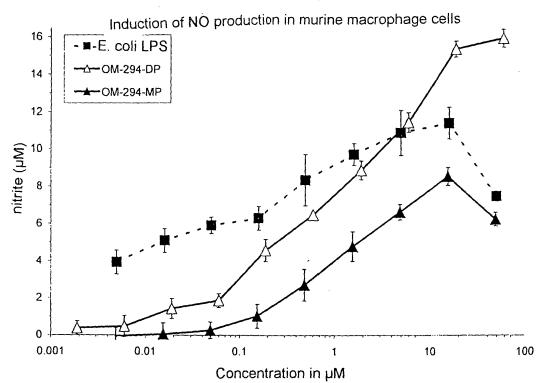


FIGURE 3

Inc > of Dextran-FITC conjugate

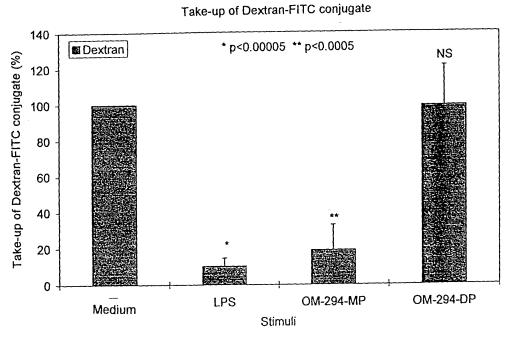


FIGURE 4

Dextran-FITC conjugate take-up: Dose related effect at low concentrations

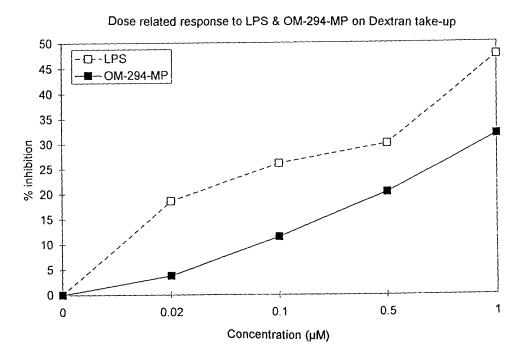


FIGURE 5

Dose related effet in terms of Dextran-FITC conjugate take-up at high concentrations

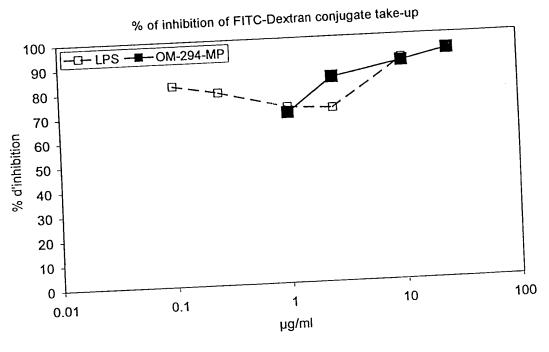
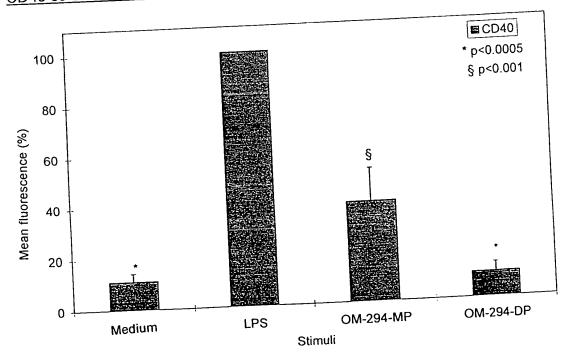


FIGURE 6

CD40 co-stimulating surface marker expression



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FIGURE 7

Expression of CD86 co-stimulating surface marker

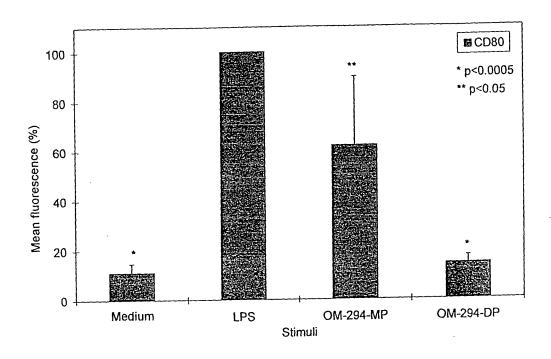
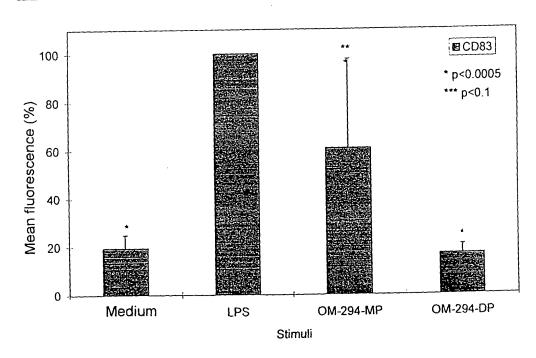


FIGURE 8

Expression of CD83 co-stimulating surface marker



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FIGURE 9

Expression of CD80 co-stimulating surface marker

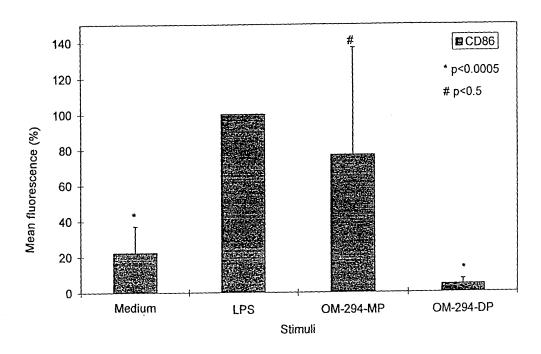
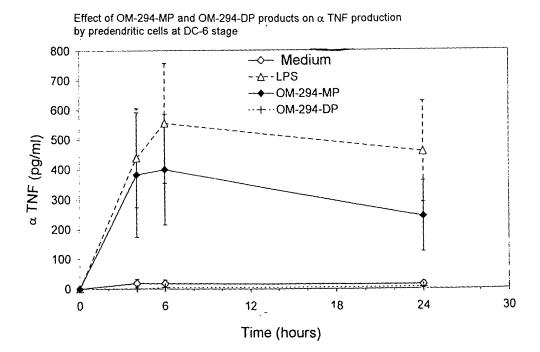


FIGURE 10 Effect of OM-294-MP and OM-294-DP products on α TNF production by predendritic cells at DC-6 stage



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FIGURE 11

Effect of OM-294-MP and OM-294-DP products on IL-12 p70 production by predendritic cells at DC-6 stage (IFN = γ IFN)

IL-12 p70 in the supernatant fluids of dendritic cells (DC-6)

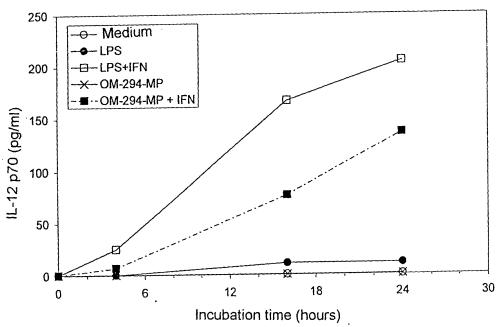
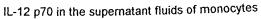


FIGURE 12

Effect of OM-294-MP products on IL-12 p70 production by monocytes

(IFN = γ IFN)



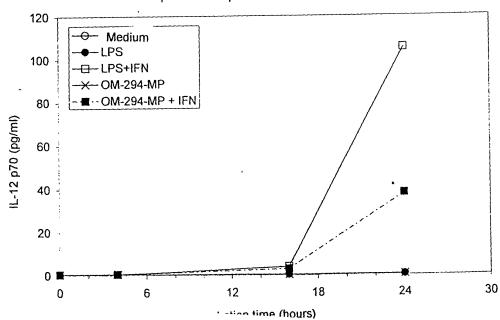


FIGURE 13

ELISA 2 after the first immunization treatment

Assay for antibodies specifically directed to Pb CS His₆ 242-310 3 weeks after the first shot (ELISA N° 2) 2.0 1.8 -A-- IFA/Pb CS 242-310, 6 His - OM-294-MP/Pb CS 242-310,6 His 1.6 Absorbance at 405 nm → OM-294-DP/Pb CS 242-310, 6 His 1.4 - X-- OM-294-MP --+--OM-294-DP 1.2 1.0 8.0 0.6 0.4 0.2 0.0 1.E-02 1.E-01 1.E-03 1.E-04 1.E-05 1.E-06

FIGURE 14

ELISA 3 after the second immunization treatment

Assay for antibodies specifically directed to Pb CS His₆ 242-310 4 weeks after the second shot (ELISA N° 3)

Serum dilution

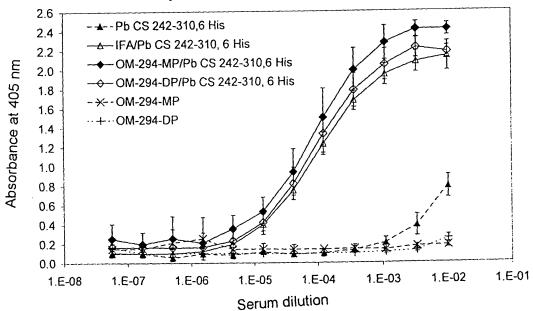
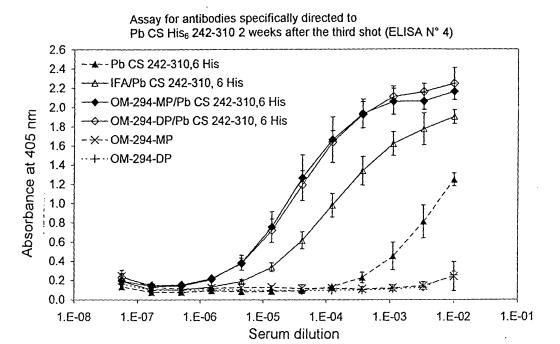


FIGURE 15
ELISA 4 after the third immunization treatment



Antibody titer before and after one, two and three immunizations treatments

Change in titer of antibodies spécifically directed to Pb CS His $_{\!6}$ 242-310 after 1, 2 and 3 injections

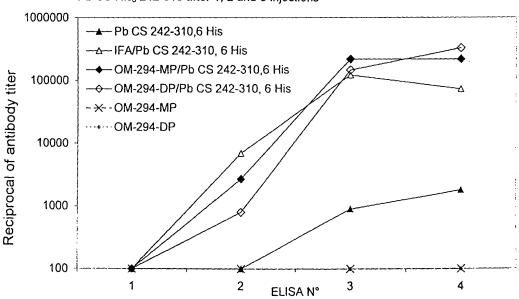


FIGURE 17

ELISPOT γ IFN-producing lymphocytes of inguinal lymph nodes stimulated by Pb CS

245 - 252 one week after the second immunization treatment

ELISPOT γ IFN-producing lymphocytes of inguinal lymph nodes one week after the second immunization treatment

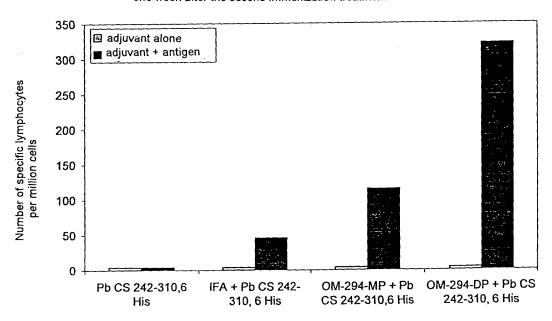


FIGURE 18 ELISPOT γ IFN-producing lymphocytes of the spleen stimulated by Pb CS 245 - 252 one week after the second immunization

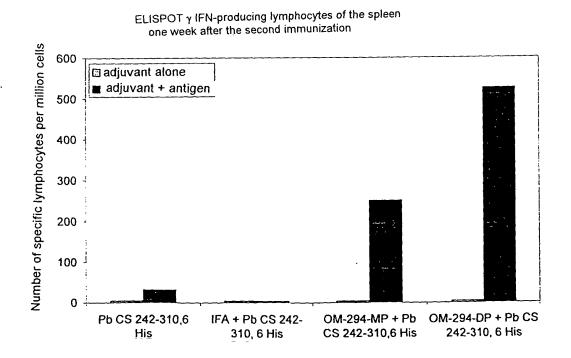


FIGURE 19
ELISPOT γ IFN-producing lymphocytes of the spleen stimulated by Pb CS 245 - 252
one week after the second immunization

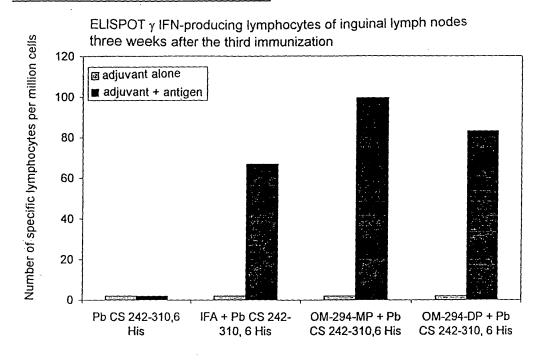


FIGURE 20

ELISPOT γ IFN-producing lymphocytes of the spleen stimulated by Pb CS 245 - 252 three weeks after the third immunization

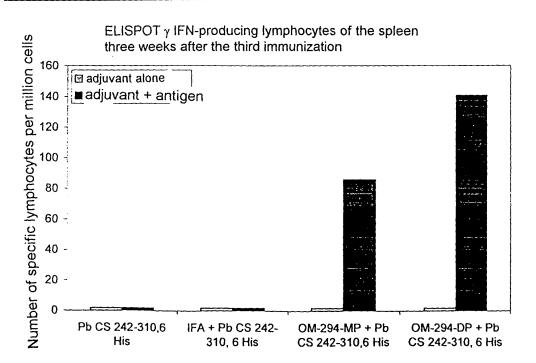


FIGURE 21

Electrophoretogram of OM-294-DP alone, of Pb CS His6-242-310 antigen alone and of Pb CS His6-242-310 - OM-294-DP complex

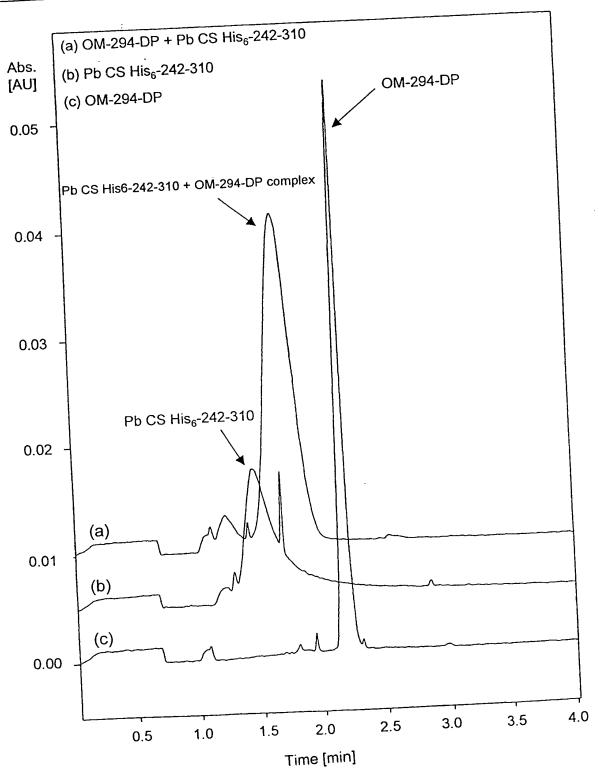


FIGURE 22

IgG1 specific antibodies directed to H1N1

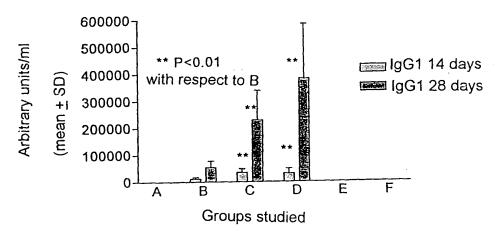


FIGURE 23

IgG2a specific antibodies directed to H1N1

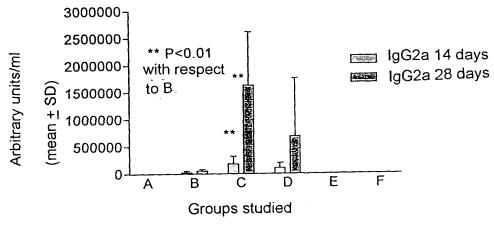


FIGURE 24

IgM specific antibodies directed to H1N1

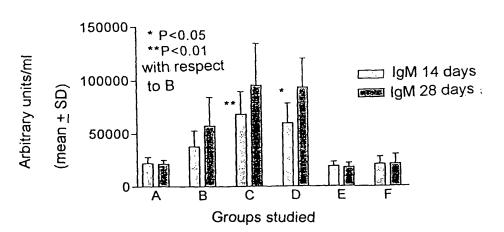




FIGURE 25

IgG1 specific antibodies directed to ovalbumin

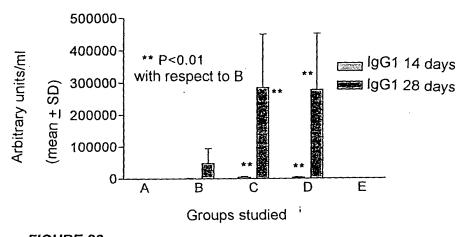


FIGURE 26

<u>IgG2a specific antibodies directed to ovalbumin</u>

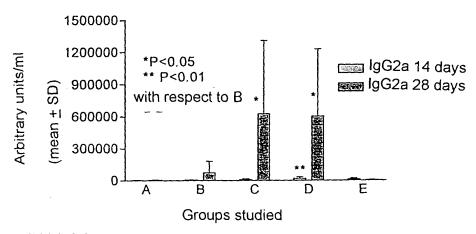
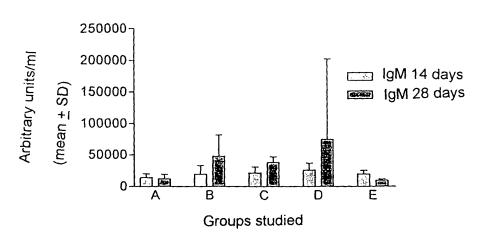


FIGURE 27

IgM specific antibodies directed to ovalbumin



IgG1 specific antibodies directed to TT

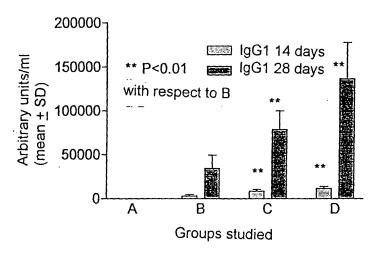
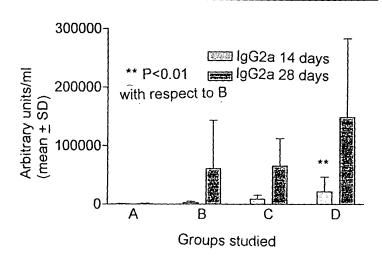


FIGURE 29

IgG2a specific antibodies directed to TT



Increase in the anti-gp63 immune response under the effect of OM-294-MP adjuvant: Comparison with BCG

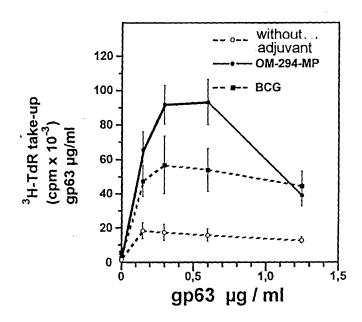
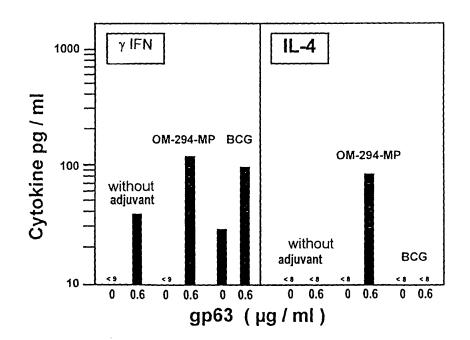


FIGURE 30 (b)



In vitro lymph node lymphocyte response derived from mice previously immunized in vivo with LmCPb antigen: effect of OM-294-MP adjuvant during the primary response

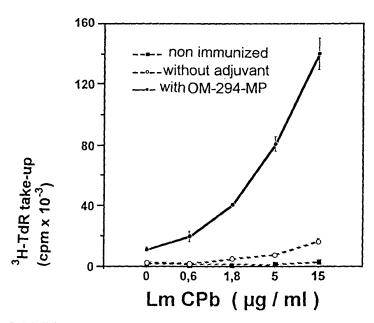


FIGURE 31 (b)

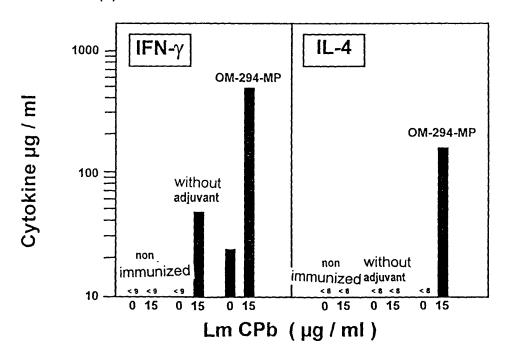




FIGURE 32 (a)

Increase in the anti-LmCPb immune response under the effect of OM-294-MP adjuvant: Comparison with BCG

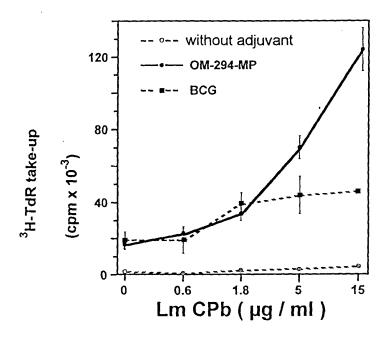
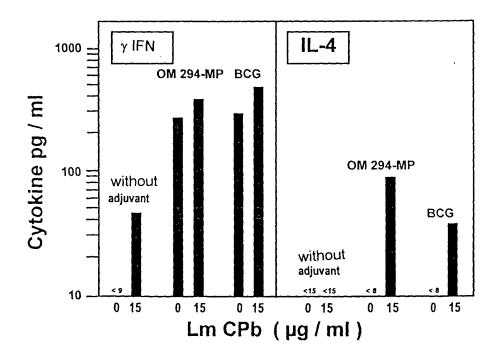
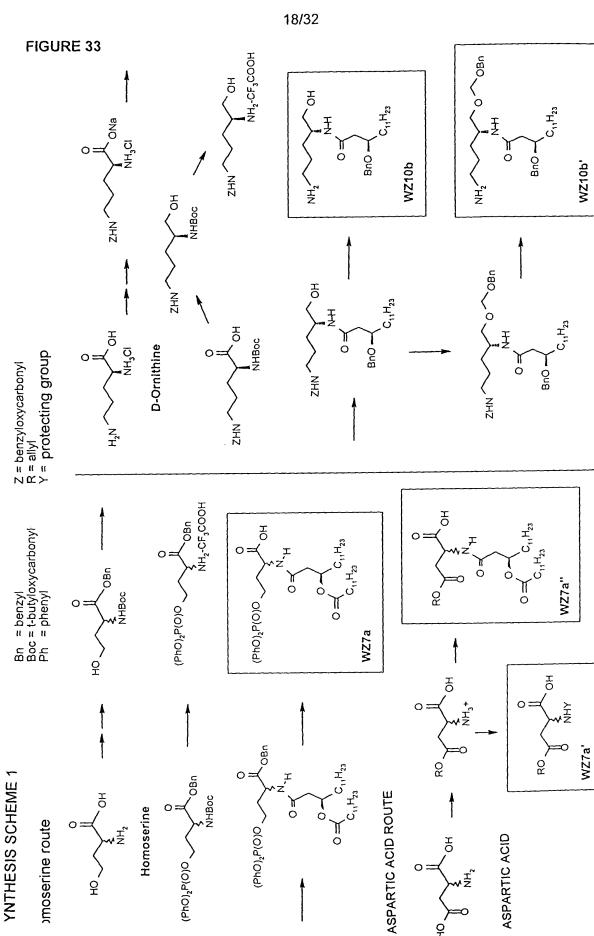


FIGURE 32 (b)





SYNTHESIS SCHEME 2

Bn ≃ benzyl Ph = phényl

$$(HO)_{2}P(O)O \longrightarrow \begin{picture}(100) \put(0.5){\line(1,0){100}} \put(0.5)$$

FIGURE 35

SYNTHESIS SCHEME 3 deprotection C11H23 0€ 10 deprotection BnO 1

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FIGURE 36

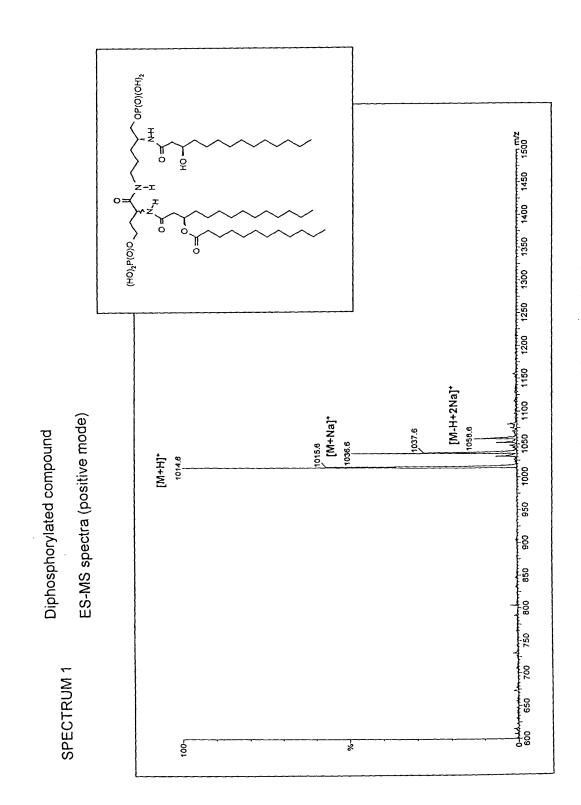
`OP(O)(OBn)₂ deprotection Bn = benzył. R = allył. Y = protecting group SYNTHESIS SCHEME 4 WZ10b BnO -WZ7aʻ



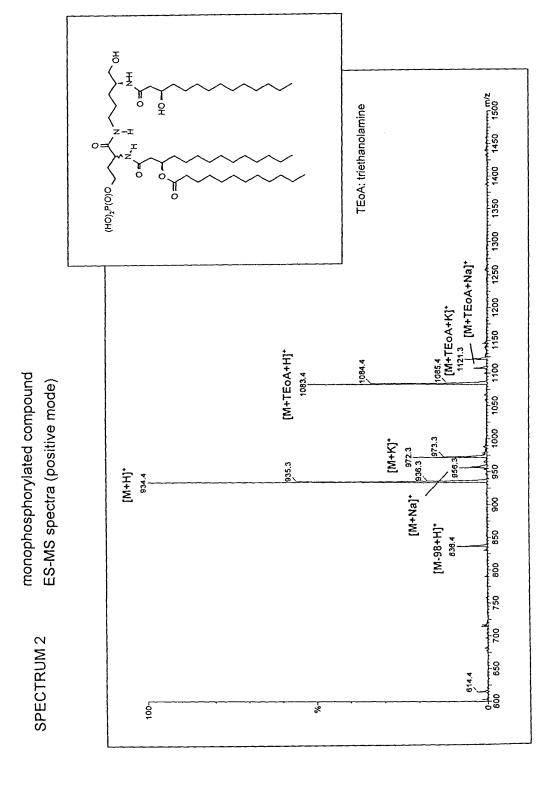
FIGURE 37

deprotection 0 0= о Д 0 (BnO)₂P(O)O Bn = benzyl R = allyl Y = protecting group SYNTHESIS SCHEME 5 WZ10b' BnO -BnO I WZ7a'

FIGURE 39



Instrumentation: Micromass Quatro II (Z-spray), triple stage quadrupole



Instrumentation: Micromass Quatro II (Z-spray), triple stage quadrupole



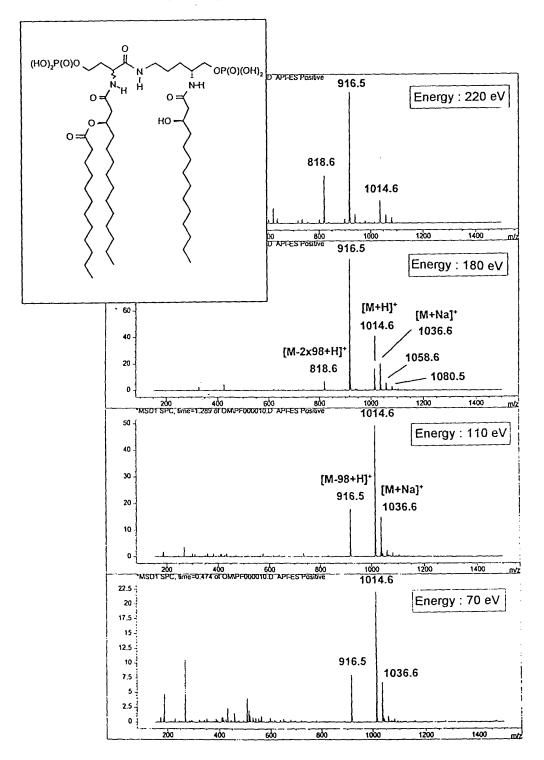


FIGURE 41

SPECTRUM 3

Diphosphorylated compound

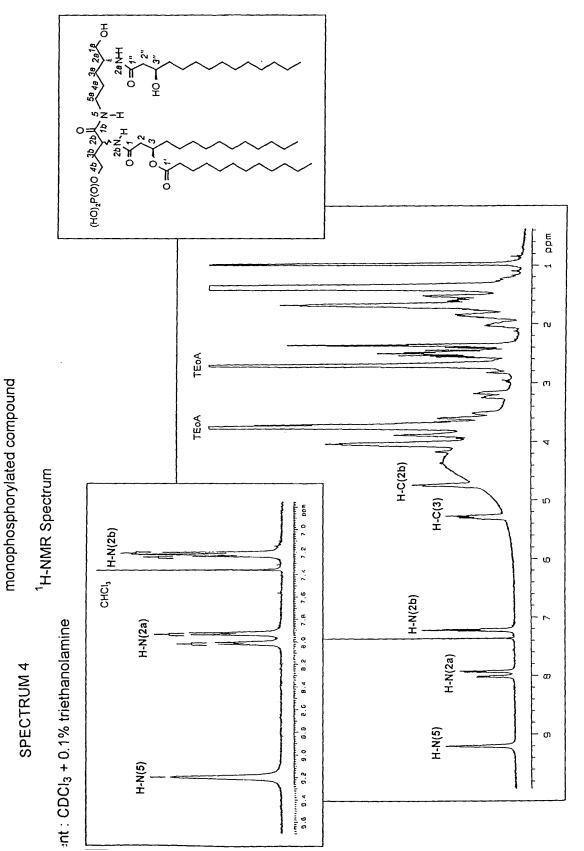
ES-MS spectra (positive mode fragmentation)



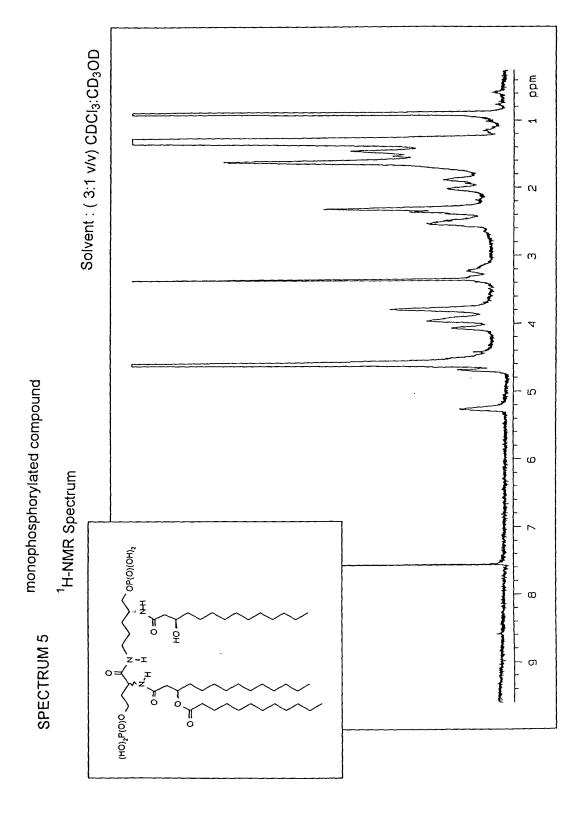
Instrumentation: Hewlett-Packard MSD, singl quadrupol



FIGURE 42



Instrumentation: Varian Unity INOVA 500 MHz



Instrumentation: Varian Unity INOVA 500 MHz

monophosphorylated compound

¹³C-NMR Spectrum

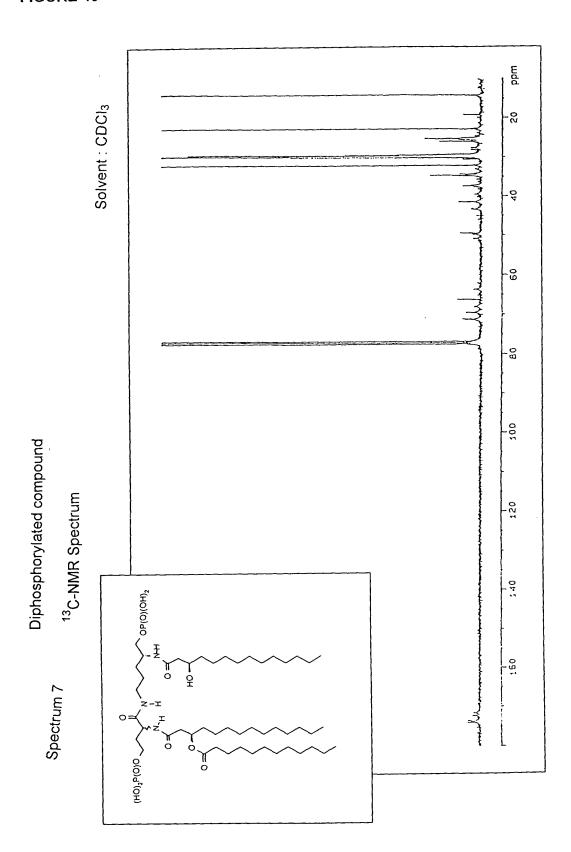
mdd Solvent: CDCI3 - ရ Q -0 . B 700 120

Instrumentation: Bruker DPX 250 MHz

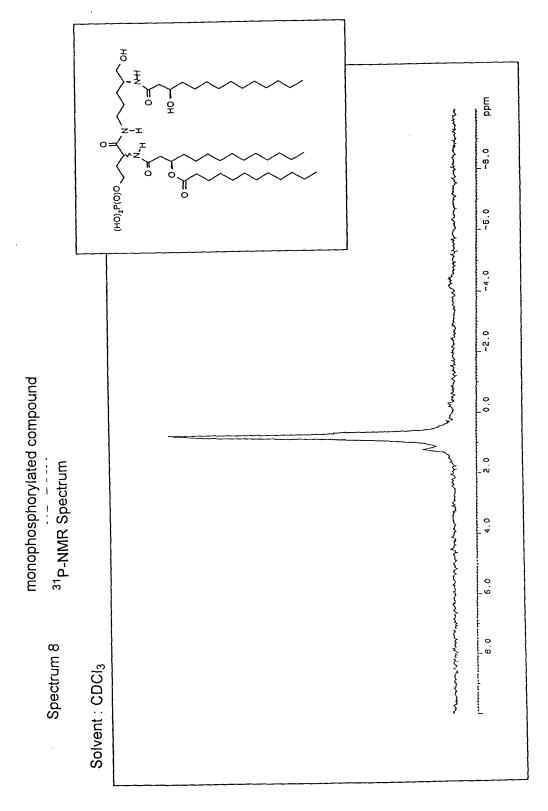
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FIGURE 45



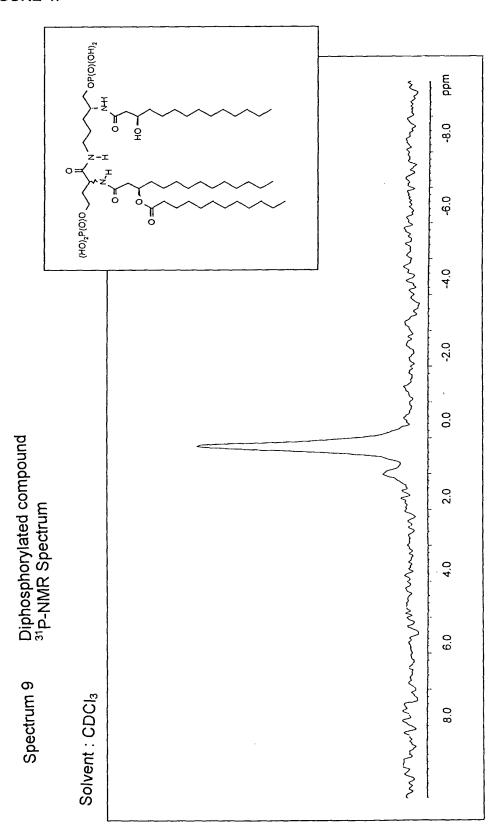
Instrumentation: Bruker DPX 250 MHz



Instrumentation: Bruker DPX 300 MHz

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FIGURE 47



Instrumentation: Bruker DPX 300 MHz